

**Amendments to the Claims**

This listing of claims will replace all prior versions and listings of claims in the application.

**Listing of Claims**

1. (Original): A method of preventing organ ischemia or reperfusion injury comprising administrating to a living subject in need thereof a pharmaceutical composition comprising:
  - a. a serine protease inhibitor; and
  - b. adenosine, an adenosine agonist or a pharmaceutically acceptable derivative or prodrug or metabolite thereof.
2. (Currently amended): The method of claim 1, wherein the serine protease inhibitor is selected from the group consisting of 4-(2-aminoethyl) benzenesulfonylfluoride, ~~□-amino-n-caproic acid~~~~-amino-n-caproic acid~~, ~~□<sub>1</sub>-antichymotrypsin~~~~α<sub>1</sub>-antichymotrypsin~~, antipain, antithrombin III, ~~□<sub>1</sub>-antitrypsin~~~~α<sub>1</sub>-antitrypsin~~, *p*-amidinophenylmethyl sulfonyl fluoride, aprotinin, cathepsin/subtilisin inhibitor (Boc-Val-Phe-NHO-Bz-*p*Cl), chymostatin (~~[(S)-1-carboxy-2-phenylethyl]-carbamoyl-□-[2-amidohexahydro-4(S)-pyrimidyl]-*(S*)-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal~~)~~[(*(S*)-1-carboxy-2-phenylethyl]-carbamoyl-α-[2-amidohexahydro-4(S)-pyrimidyl]-(*S*)-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal~~), chymotrypsin inhibitor I, 3,4-dichloroisocoumarin, diisopropylfluoro phosphate, dipeptidylpeptidase IV inhibitor I (Ile-Pro-Ile), dipeptidylpeptidase IV inhibitor II (H-Glu-(NHO-Bz)-Pyr), ecotin, elastase inhibitor I (Boc-Ala-Ala-Ala-NHO-Bz), ~~□<sub>2</sub>-macroglobulin~~~~α<sub>2</sub>-macroglobulin~~, PPACK (*D*-Phe-Pro-Arg-chloromethylketone), PPACK II, *N*<sup>a</sup>-tosyl-Lys chloromethyl ketone, *N*<sup>a</sup>-tosyl-Phe chloromethyl ketone, and any mixture thereof.
3. (Original): The method of claim 1, wherein the adenosine agonist or pharmaceutically

acceptable derivative is selected from the group consisting of AB-MECA ( $N^6$ -4-aminobenzyl-5'-N-methyl carboxamidoadenosine), CPA ( $N^6$ -cyclopentyladenosine), ADAC ( $N^6$ - [4-[[[4-[[[(2-aminoethyl) amino] carbonyl] methyl]-anilino] carbonyl] methyl] phenyl] adenosine), CCPA (2-chloro- $N^6$ -cyclopentyladenosine), CHA ( $N^6$ -cyclohexyladenosine), GR79236 ( $N^6$ -[1*S*, *trans*,2-hydroxy cyclopentyl] adenosine), S-ENBA ((2*S*)-  $N^6$ -(2-endonorbanyl)adenosine), IAB-MECA ( $N^6$ -(4-amino-3-iodobenzyl)adenosine-5'-N-methylcarboxamidoadenosine), *R*-PIA (*R*- $N^6$ -(phenyl isopropyl) adenosine), ATL146e (4-{3-[6-amino-9-(5-ethylcarbamoyl-3,4-dihydroxy-tetrahydro-furan-2-yl)-9H-purin-2-yl]-prop-2-ynyl}-cyclohexanecarboxylic acid methyl ester), CGS-21680 (APEC or 2-[*p*-(2-carbonyl-ethyl)-phenyl ethyl amino]-5'-N-ethylcarboxamidoadenosine), CV1808 (2-phenylaminoadenosine), HENECA (2-hex-1-ynyl-5'-N-ethylcarboxamido adenosine), NECA (5'-N-ethyl-carboxamido adenosine), PAPA-APEC (2-(4-[2-[(4-aminophenyl) methyl carbonyl] ethyl] phenyl) ethylamino-5'-N-ethyl carboxamidoadenosine), DITC-APEC (2-[*p*-(4-isothiocyanatophenylamino thiocarbonyl-2-ethyl)-phenylethylamino]-5'-N-ethylcarboxamido adenosine), DPMA ( $N^6$ -(2(3,5-dimethoxy phenyl)-2-(2-methyl phenyl) ethyl)adenosine), S-PHPNECA ((*S*)-2-phenylhydroxypropynyl-5'-N-ethylcarbox amidoadenosine), WRC-0470 (2-cyclohexylmethylenedihydrazinoadenosine), AMP-579 (1*S*-[1a,2b,3b,4a(*S*\*)]]-4-[7-[[2-(3-chloro-2-thienyl)-1-methylpropyl]amino]-3H-imidazo [4,5-b] pyridyl-3-yl] cyclopentane carboxamide), IB-MECA ( $N^6$ - (3-iodobenzyl) adenosine -5'-N-methyluronamide), 2-CIADO (2-chloroadenosine), I-ABA ( $N^6$ -(4-amino-3-iodobenzyl)adenosine), *S*-PIA (*S*- $N^6$ -(phenylisopropyl) adenosine), 2-[(2-aminoethyl-aminocarbonylethyl) phenylethyl amino]-5'-N-ethyl-carboxamido adenosine, 2-Cl-IB-MECA (2-chloro- $N^6$ - (3-iodobenzyl)adenosine-5'-N-methyluronamide), polyadenylic acid, and any mixture thereof.

4. (Original): A pharmaceutical composition comprising:
  - a. a serine protease inhibitor; and
  - b. adenosine, an adenosine agonist or a pharmaceutically acceptable derivative or prodrug or metabolite thereof.

5. (Currently amended): The pharmaceutical composition of claim 4, wherein the serine protease inhibitor is selected from the group consisting of 4-(2-aminoethyl) benzenesulfonylfluoride, ~~□-amino-n-caproic-acide-amino-n-caproic acid, □-1-antichymotrypsinα<sub>1</sub>-antichymotrypsin~~, antipain, antithrombin III, ~~□-1-antitrypsinα<sub>1</sub>-antitrypsin~~, ~~p-amidino phenylmethylsulfonyl fluoride, aprotinin, cathepsin/subtilisin inhibitor (Boc-Val-Phe-NHO-Bz-pCl), chymostatin ([(S)-1-carboxy-2-phenylethyl]-carbamoyl-□-[2-amidohexa hydro-4(S)-pyrimidyl]- (S)-glycyl-[A = Leu, B = Val, or C = Ile]-phenyl alaninal) [(S)-1-carboxy-2-phenylethyl]-carbamoyl-α- [2-amidohexa hydro-4(S)-pyrimidyl]- (S)-glycyl-[A = Leu, B = Val, or C = Ile]-phenyl alaninal~~, chymotrypsin inhibitor I, 3,4-dichloroisocoumarin, diisopropylfluorophosphate, dipeptidylpeptidase IV inhibitor I (Ile-Pro-Ile), dipeptidylpeptidase IV inhibitor II (H-Glu-(NHO-Bz)-Pyr), ecotin, elastase inhibitor I (Boc-Ala-Ala-Ala-NHO-Bz), ~~□<sub>2</sub>-macroglobulinα<sub>2</sub>-macroglobulin, PPACK (D-Phe-Pro-Arg-chloromethylketone), PPACK II, N<sup>a</sup>-tosyl-Lys chloromethyl ketone, N<sup>a</sup>-tosyl-Phe chloromethyl ketone, and any mixture thereof.~~

6. (Original): The pharmaceutical composition of claim 4, wherein the adenosine agonist or pharmaceutically acceptable derivative is selected from the group consisting of AB-MECA (*N*<sup>6</sup>-4-aminobenzyl-5'-*N*-methylcarboxamido adenosine), CPA (*N*<sup>6</sup>-cyclopentyl adenosine), ADAC (*N*<sup>6</sup>-[4-[[4-[[[(2-aminoethyl) amino] carbonyl] methyl]-anilino] carbonyl] methyl] phenyl adenosine), CCPA (2-chloro-*N*<sup>6</sup>-cyclopentyl adenosine), CHA (*N*<sup>6</sup>-cyclohexyl adenosine), GR79236 (*N*<sup>6</sup>-[1*S*, *trans*,2-hydroxy cyclopentyl] adenosine), *S*-ENBA ((2*S*)-*N*<sup>6</sup>-(2-endonorbanyl) adenosine), IAB-MECA (*N*<sup>6</sup>-(4-amino-3-iodobenzyl) adenosine-5'-*N*-methylcarboxamido adenosine), *R*-PIA (*R*-*N*<sup>6</sup>-(phenylisopropyl) adenosine), ATL146e (4-{3-[6-amino-9-(5-ethyl carbamoyl)-3,4-dihydroxy-tetrahydro-furan-2-yl]-9*H*-purin-2-yl]-prop-2-ynyl}-cyclohexane carboxylic acid methyl ester), CGS-21680 (APEC or 2-[*p*-(2-carbonyl-ethyl)-phenyl ethyl amino]-5'-*N*-ethylcarboxamido adenosine), CV1808 (2-phenylamino adenosine), HENECA (2-hex-1-ynyl-5'-*N*-ethylcarboxamido adenosine), NECA (5'-*N*-ethyl-carboxamido adenosine), PAPA-APEC (2-(4-[2-[(4-aminophenyl)methylcarbonyl]ethyl] phenyl) ethylamino-5'-*N*-

ethyl carboxamidoadenosine), DITC-APEC (2-[*p*-(4-isothiocyanatophenylamino thiocarbonyl-2-ethyl)-phenylethylamino]-5'-*N*-ethylcarboxamidoadenosine), DPMA (*N*<sup>6</sup>-(2(3,5-dimethoxy phenyl)-2-(2-methylphenyl)ethyl)adenosine), *S*-PHPNECA ((*S*)-2-phenylhydroxypropynyl-5'-*N*-ethylcarboxamidoadenosine), WRC-0470 (2-cyclohexyl methylidenehydrazinoadenosine), AMP-579 (1*S*-[1a,2b,3b,4a(S\*)]-4-[7-[[2-(3-chloro-2-thienyl)-1-methylpropyl]amino]-3H-imidazo [4,5-b] pyridyl-3-yl] cyclopentane carboxamide), IB-MECA (*N*<sup>6</sup>-(3-iodobenzyl)adenosine-5'-*N*-methyluronamide), 2-CIADO (2-chloroadenosine), I-ABA (*N*<sup>6</sup>-(4-amino-3-iodobenzyl) adenosine), *S*-PIA (*S*-*N*<sup>6</sup>-(phenylisopropyl)adenosine), 2-[(2-aminoethyl-aminocarbonylethyl) phenylethyl amino]-5'-*N*-ethyl-carboxamidoadenosine, 2-Cl-IB-MECA (2-chloro-*N*<sup>6</sup>-(3-iodobenzyl) adenosine-5'-*N*-methyluronamide), polyadenylic acid, and any mixture thereof.

7. (Original): A method of preventing organ ischemia or reperfusion injury comprising concomitantly administering to a living subject in need thereof
  - a. a serine protease inhibitor; and
  - b. adenosine, an adenosine agonist or a pharmaceutically acceptable derivative or prodrug or metabolite thereof.
8. (Currently amended): The method of claim 7, wherein the serine protease inhibitor is selected from the group consisting of 4-(2-aminoethyl) benzenesulfonylfluoride,  $\square$ -amino-*n*-caproic acid-amino-*n*-caproic acid,  $\square$ <sub>4</sub>-antichymotrypsin $\alpha$ <sub>1</sub>-antichymotrypsin, antipain, antithrombin III,  $\square$ <sub>4</sub>-antitrypsin $\alpha$ <sub>1</sub>-antitrypsin, *p*-amidinophenylmethyl sulfonyl fluoride, aprotinin, cathepsin/subtilisin inhibitor (Boc-Val-Phe-NHO-Bz-*p*Cl), chymostatin (~~([(S)-1-carboxy-2-phenylethyl]-carbamoyl- $\square$ -[2-amidohexahydro-4-(*S*)-pyrimidyl]-(*S*)-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal)~~~~([(S)-1-carboxy-2-phenylethyl]-carbamoyl- $\alpha$ -[2-amidohexahydro-4-(*S*)-pyrimidyl]-(*S*)-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal)~~, chymotrypsin inhibitor I, 3,4-dichloroisocoumarin, diisopropylfluorophosphate, dipeptidylpeptidase IV inhibitor I (Ile-Pro-Ile), dipeptidylpeptidase IV inhibitor II (H-Glu-(NHO-Bz)-Pyr), ecotin, elastase inhibitor I (Boc-Ala-Ala-Ala-NHO-Bz),  $\square$ <sub>2</sub>-macroglobulin $\alpha$ <sub>2</sub>-macroglobulin, PPACK (*D*-Phe-Pro-

Arg-chloromethylketone), PPACK II, *N*<sup>a</sup>-tosyl-Lys chloromethyl ketone, *N*<sup>a</sup>-tosyl-Phe chloromethyl ketone, and any mixture thereof.

9. (Original): The method of claim 7, wherein the adenosine agonist or pharmaceutically acceptable derivative is selected from the group consisting of AB-MECA (*N*<sup>6</sup>-4-aminobenzyl-5'-*N*-methylcarboxamidoadenosine), CPA (*N*<sup>6</sup>-cyclopentyladenosine), ADAC (*N*<sup>6</sup>-[4-[[[4-[[[(2-aminoethyl) amino] carbonyl] methyl]-anilino] carbonyl] methyl] phenyl] adenosine), CCPA (2-chloro-*N*<sup>6</sup>-cyclopentyladenosine), CHA (*N*<sup>6</sup>-cyclohexyladenosine), GR79236 (*N*<sup>6</sup>-[1*S*, *trans*,2-hydroxycyclopentyl] adenosine), *S*-ENBA ((2*S*)- *N*<sup>6</sup>-(2-endonorbanyl)adenosine), IAB-MECA (*N*<sup>6</sup>-(4-amino-3-iodobenzyl)adenosine-5'-*N*-methylcarboxamidoadenosine), *R*-PIA (*R*-*N*<sup>6</sup>-(phenylisopropyl) adenosine), ATL146e (4-{3-[6-amino-9-(5-ethylcarbamoyl-3,4-dihydroxy-tetrahydro-furan-2-yl)-9H-purin-2-yl]-prop-2-ynyl}-cyclohexanecarboxylic acid methyl ester), CGS-21680 (APEC or 2-[*p*-(2-carbonyl-ethyl)-phenyl ethyl amino]-5'-*N*-ethylcarboxamido adenosine), CV1808 (2-phenylaminoadenosine), HENECA (2-hex-1-ynyl-5'-*N*-ethylcarboxamido adenosine), NECA (5'-*N*-ethyl-carboxamido adenosine), PAPA-APEC (2-(4-[2-[(4-amino phenyl) methylcarbonyl]ethyl] phenyl) ethylamino-5'-*N*-ethyl carboxamidoadenosine), DITC-APEC (2-[*p*-(4-isothiocyanatophenylamino thiocarbonyl-2-ethyl)-phenylethylamino]-5'-*N*-ethylcarboxamidoadenosine), DPMA (*N*<sup>6</sup>-(2(3,5-dimethoxy phenyl)-2-(2-methylphenyl)ethyl)adenosine), *S*-PHPNECA ((*S*)-2-phenylhydroxypropynyl-5'-*N*-ethylcarboxamidoadenosine), WRC-0470 (2-cyclohexylmethylenedihydrazinoadenosine), AMP-579 (1*S*-[1a,2b,3b,4a(*S*\*)])-4-[7-[[2-(3-chloro-2-thienyl)-1-methylpropyl]amino]-3H-imidazo [4,5-b] pyridyl-3-yl] cyclopentane carboxamide), IB-MECA (*N*<sup>6</sup>- (3-iodobenzyl)adenosine-5'-*N*-methyluronamide), 2-CIADO (2-chloroadenosine), I-ABA (*N*<sup>6</sup>-(4-amino-3-iodobenzyl) adenosine), *S*-PIA (*S*-*N*<sup>6</sup>-(phenylisopropyl)adenosine), 2-[(2-aminoethyl-aminocarbonylethyl) phenylethyl amino]-5'-*N*-ethyl-carboxamidoadenosine, 2-Cl-IB-MECA (2-chloro-*N*<sup>6</sup>- (3-iodobenzyl)adenosine-5'-*N*-methyluronamide), polyadenylic acid, and any mixture thereof.

10. (Original): A method of preventing organ ischemia or reperfusion injury comprising administering to a living subject in need thereof sequentially in any order
  - a. a serine protease inhibitor; and
  - b. adenosine, an adenosine agonist or a pharmaceutically acceptable derivative or prodrug or metabolite thereof.
11. (Currently amended): The method of claim 10, wherein the serine protease inhibitor is selected from the group consisting of 4-(2-aminoethyl) benzenesulfonylfluoride, ~~amino-n-caproic acid~~ ~~amino-n-caproic acid~~, ~~α<sub>1</sub>-antichymotrypsin~~<sub>α<sub>1</sub></sub>-antichymotrypsin, antipain, antithrombin III, ~~α<sub>1</sub>-antitrypsin~~<sub>α<sub>1</sub></sub>-antitrypsin, *p*-amidinophenylmethyl sulfonyl fluoride, aprotinin, cathepsin/subtilisin inhibitor (Boc-Val-Phe-NHO-Bz-*p*Cl), chymostatin (~~([(S)-1-carboxy-2-phenylethyl]-carbamoyl-α-[2-amidohexahydro-4(S)-pyrimidyl]-S-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal)~~<sub>[(S)-1-carboxy-2-phenylethyl]-carbamoyl-α-[2-amidohexahydro-4(S)-pyrimidyl]-S-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal</sub>), chymotrypsin inhibitor I, 3,4-dichloroisocoumarin, diisopropylfluorophosphate, dipeptidylpeptidase IV inhibitor I (Ile-Pro-Ile), dipeptidylpeptidase IV inhibitor II (H-Glu-(NHO-Bz)-Pyr), ecotin, elastase inhibitor I (Boc-Ala-Ala-Ala-NHO-Bz), ~~α<sub>2</sub>-macroglobulin~~<sub>α<sub>2</sub></sub>-macroglobulin, PPACK (*D*-Phe-Pro-Arg-chloromethylketone), PPACK II, *N*<sup>2</sup>-tosyl-Lys chloromethyl ketone, *N*<sup>2</sup>-tosyl-Phe chloromethyl ketone, and any mixture thereof.
12. (Original): The method of claim 10, wherein the adenosine agonist or pharmaceutically acceptable derivative is selected from the group consisting of AB-MECA (*N*<sup>6</sup>-4-aminobenzyl-5'-*N*-methylcarboxamidoadenosine), CPA (*N*<sup>6</sup>-cyclopentyladenosine), ADAC (*N*<sup>6</sup>-[4-[[4-[[[(2-aminoethyl) amino] carbonyl] methyl]-anilino] carbonyl] methyl] phenyl] adenine), CCPA (2-chloro-*N*<sup>6</sup>-cyclopentyladenosine), CHA (*N*<sup>6</sup>-cyclohexyladenosine), GR79236 (*N*<sup>6</sup>-[1*S*, *trans*,2-hydroxycyclopentyl] adenine), *S*-ENBA ((2*S*)-*N*<sup>6</sup>-(2-endonorbanyl)adenosine), IAB-MECA (*N*<sup>6</sup>-(4-amino-3-iodobenzyl)adenosine-5'-*N*-methylcarboxamidoadenosine), *R*-PIA (*R*-*N*<sup>6</sup>-(phenylisopropyl) adenosine), ATL146e (4-{3-[6-amino-9-(5-ethylcarbamoyl-3,4-

dihydroxy-tetrahydro-furan-2-yl)-9H-purin-2-yl]-prop-2-ynyl}-cyclohexanecarboxylic acid methyl ester), CGS-21680 (APEC or 2-[*p*-(2-carbonyl-ethyl)-phenyl ethyl amino]-5'-*N*-ethylcarbox amido adenosine), CV1808 (2-phenylaminoadenosine), HENECA (2-hex-1-ynyl-5'-*N*-ethylcarboxamido adenosine), NECA (5'-*N*-ethyl-carboxamido adenosine), PAPA-APEC (2-(4-[2-[(4-amino phenyl) methylcarbonyl]ethyl] phenyl) ethylamino-5'-*N*-ethyl carboxamidoadenosine), DITC-APEC (2-[*p*-(4-isothiocyanatophenylamino thiocarbonyl-2-ethyl)-phenylethylamino]-5'-*N*-ethylcarboxamidoadenosine), DPMA (*N*<sup>6</sup>-(2(3,5-dimethoxy phenyl)-2-(2-methylphenyl)ethyl) adenosine), *S*-PHPNECA ((*S*)-2-phenylhydroxypropynyl-5'-*N*-ethylcarboxamidoadenosine), WRC-0470 (2-cyclohexylmethyldenehydrazinoadenosine), AMP-579 (1*S*-[1a,2b,3b,4a(*S*\*)]-4-[7-[[2-(3-chloro-2-thienyl)-1-methylpropyl]amino]-3H-imidazo [4,5-b] pyridyl-3-yl] cyclo pentane carboxamide), IB-MECA (*N*<sup>6</sup>- (3-iodobenzyl)adenosine-5'-*N*-methyluronamide), 2-CIADO (2-chloroadenosine), I-ABA (*N*<sup>6</sup>-(4-amino-3-iodobenzyl) adenosine), *S*-PIA (*S*-*N*<sup>6</sup>-(phenylisopropyl)adenosine), 2-[(2-aminoethyl-aminocarbonylethyl) phenylethyl amino]-5'-*N*-ethyl-carboxamidoadenosine, 2-Cl-IB-MECA (2-chloro-*N*<sup>6</sup>- (3-iodobenzyl)adenosine-5'-*N*-methyluronamide), polyadenylic acid, and any mixture thereof.

13. (Original): A method of preventing organ or tissue injury at a predetermined point or period of intervention comprising administrating to a living subject in need thereof a pharmaceutical composition comprising:
  - a. a serine protease inhibitor; and
  - b. adenosine, an adenosine agonist or a pharmaceutically acceptable derivative or prodrug or metabolite thereof.
14. (Original): The method of claim 13, wherein the organ or tissue injury is related to at least one of cardiac surgery, non-surgical cardiac revascularization, organ transplantation, perfusion, ischemia, reperfusion, ischemia-reperfusion injury, oxidant injury, cytokine induced injury, shock induced injury, resuscitations injury, and apoptosis.

15. (Original): The method of claim 13, wherein the administrating is taken at the predetermined point of intervention related to at least one of pre-treatment regimen, pharmacological preconditioning, reperfusion, or post interventional therapy, wherein the pharmacological preconditioning is a treatment administered before the ischemic intervention followed by a brief period of reperfusion or washout.
16. (Currently amended): The method of claim 13, wherein the serine protease inhibitor is selected from the group consisting of 4-(2-aminoethyl) benzenesulfonylfluoride, ~~□-amino-n-caproic acid~~ ~~□-amino-n-caproic acid~~, ~~□<sub>1</sub>-antichymotrypsin~~ ~~□<sub>1</sub>-antichymotrypsin~~, antipain, antithrombin III, ~~□<sub>1</sub>-antitrypsin~~ ~~□<sub>1</sub>-antitrypsin~~, *p*-amidinophenylmethyl sulfonyl fluoride, aprotinin, cathepsin/subtilisin inhibitor (Boc-Val-Phe-NHO-Bz-*p*Cl), chymostatin (~~[(S)-1-carboxy-2-phenylethyl] carbamoyl~~ ~~□~~ [2-amidohexahydro-4(S)-pyrimidyl]-(S)-glycyl [A = Leu, B = Val, or C = Ile] phenylalaninal) (~~[(S)-1-carboxy-2-phenylethyl] carbamoyl~~- $\alpha$ - [2-amidohexahydro-4(S)-pyrimidyl]-(S)-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal), chymotrypsin inhibitor I, 3,4-dichloroisocoumarin, diisopropylfluorophosphate, dipeptidylpeptidase IV inhibitor I (Ile-Pro-Ile), dipeptidylpeptidase IV inhibitor II (H-Glu-(NHO-Bz)-Pyr), ecotin, elastase inhibitor I (Boc-Ala-Ala-Ala-NHO-Bz), ~~□<sub>2</sub>-macroglobulin~~ ~~□<sub>2</sub>-macroglobulin~~, PPACK (*D*-Phe-Pro-Arg-chloromethylketone), PPACK II, *N*<sup>a</sup>-tosyl-Lys chloromethyl ketone, *N*<sup>a</sup>-tosyl-Phe chloromethyl ketone, and any mixture thereof.
17. (Original): The method of claim 13, wherein the adenosine agonist or pharmaceutically acceptable derivative is selected from the group consisting of AB-MECA (*N*<sup>6</sup>-4-aminobenzyl-5'-*N*-methylcarboxamidoadenosine), CPA (*N*<sup>6</sup>-cyclopentyladenosine), ADAC (*N*<sup>6</sup>- [4-[[[4-[[[(2-aminoethyl) amino] carbonyl] methyl]-anilino] carbonyl] methyl] phenyl] adenine), CCPA (2-chloro-*N*<sup>6</sup>-cyclopentyladenosine), CHA (*N*<sup>6</sup>-cyclohexyladenosine), GR79236 (*N*<sup>6</sup>-[1*S*, *trans*,2-hydroxycyclopentyl] adenine), *S*-ENBA ((2*S*)- *N*<sup>6</sup>-(2-endonorbanyl)adenosine), IAB-MECA (*N*<sup>6</sup>-(4-amino-3-iodobenzyl)adenosine-5'-*N*-methylcarboxamidoadenosine), *R*-PIA (*R*-*N*<sup>6</sup>-(phenylisopropyl) adenosine), ATL146e (4-{3-[6-amino-9-(5-ethylcarbamoyl)-3,4-

dihydroxy-tetrahydro-furan-2-yl)-9H-purin-2-yl]-prop-2-ynyl}-cyclohexanecarboxylic acid methyl ester), CGS-21680 (APEC or 2-[*p*-(2-carbonyl-ethyl)-phenyl ethyl amino]-5'-*N*-ethylcarboxamidoadenosine), CV1808 (2-phenylaminoadenosine), HNECA (2-hex-1-ynyl-5'-*N*-ethylcarboxamido adenosine), NECA (5'-*N*-ethyl-carboxamido adenosine), PAPA-APEC (2-(4-[2-[(4-aminophenyl) methyl carbonyl]ethyl] phenyl) ethylamino-5'-*N*-ethyl carboxamidoadenosine), DITC-APEC (2-[*p*-(4-isothiocyanatophenylamino thiocarbonyl-2-ethyl)-phenylethylamino]-5'-*N*-ethylcarboxamidoadenosine), DPMA (*N*<sup>6</sup>-(2(3,5-dimethoxy phenyl)-2-(2-methyl phenyl) ethyl)adenosine), S-PHPNECA ((*S*)-2-phenylhydroxypropynyl-5'-*N*-ethyl carboxamidoadenosine), WRC-0470 (2-cyclohexylmethyldenehydrazinoadenosine), AMP-579 (1*S*-[1a,2b,3b,4a(*S*\*)])-4-[7-[[2-(3-chloro-2-thienyl)-1-methylpropyl]amino]-3H-imidazo [4,5-b] pyridyl-3-yl] cyclopentane carboxamide), IB-MECA (*N*<sup>6</sup>- (3-iodo benzyl)adenosine-5'-*N*-methyluronamide), 2-CIADO (2-chloroadenosine), I-ABA (*N*<sup>6</sup>-(4-amino-3-iodobenzyl)adenosine), S-PIA (*S*-*N*<sup>6</sup>-(phenylisopropyl)adenosine), 2-[(2-amino ethyl-aminocarbonylethyl) phenylethyl amino]-5'-*N*-ethyl-carboxamidoadenosine, 2-Cl-IB-MECA (2-chloro-*N*<sup>6</sup>- (3-iodobenzyl)adenosine-5'-*N*-methyluronamide), polyadenylic acid, and any mixture thereof.

18. (Original): A method of preventing organ ischemia or reperfusion injury comprising administrating to a living subject in need thereof a pharmaceutical composition comprising:
  - a. a protease inhibitor; and
  - b. an agent that alters activities of G protein coupled receptors and cAMP, an analog or a pharmaceutically acceptable derivative or prodrug or metabolite thereof.
19. (Currently amended): The method of claim 18, wherein the protease inhibitor is selected from the group consisting of 4-(2-aminoethyl) benzenesulfonylfluoride, ~~□-amino-*n*-caproic acid~~-amino-*n*-caproic acid, ~~□-antichymotrypsin~~ $\alpha_1$ -antichymotrypsin, antipain, antithrombin III, ~~□-antitrypsin~~ $\alpha_1$ -antitrypsin, *p*-amidinophenylmethyl sulfonyl fluoride, aprotinin, cathepsin/subtilisin inhibitor (Boc-Val-Phe-NHO-Bz-*p*Cl), chymostatin ([(*S*)-

~~1-carboxy-2-phenylethyl]-carbamoyl-□-[2-amidohexahydro-4(S)-pyrimidyl]- (S)-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal)([(S)-1-carboxy-2-phenylethyl]-carbamoyl- $\alpha$ -[2-amidohexahydro-4(S)-pyrimidyl]- (S)-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal), chymotrypsin inhibitor I, 3,4-dichloroisocoumarin, diisopropylfluoro phosphate, dipeptidylpeptidase IV inhibitor I (Ile-Pro-Ile), dipeptidylpeptidase IV inhibitor II (H-Glu-(NHO-Bz)-Pyr), ecotin, elastase inhibitor I (Boc-Ala-Ala-Ala-NHO-Bz),  $\square_2$ -macroglobulin $\alpha_2$ -macroglobulin, PPACK (*D*-Phe-Pro-Arg-chloromethylketone), PPACK II,  $N^{\alpha}$ -tosyl-Lys chloromethyl ketone,  $N^{\alpha}$ -tosyl-Phe chloromethyl ketone, acetyl-pepstatin (Ac-Val-Val-(3*S*,4*S*)-Sta-Ala-(3*S*,4*S*)-Sta-OH), calpain inhibitor I (*N*-acetyl-Leu-Leu-norleucinal), calpain inhibitor II (*N*-acetyl-Leu-Leu-Met-CHO), amastatin ([(2*S*, 2*R*)]-3-amino-2-hydroxy-5-methylhexanoyl] -Val-Val-Asp-OH), arphamenine A ((2*R*,5*S*)-5-amino-8-guanidino-4-oxo-2-phenylmethyl octanoic acid), arphamenine B ((2*R*,5*S*)-5-amino-8-guanidino-4-oxo-2-*p*-hydroxyphenyl methyloctanoic acid), benzamidine, bestatin ([(2*S*, 2*R*)-3-amino-2-hydroxy-4-phenyl butanoyl] -*L*-Leucine), CA-074 ((*L*-3-*trans*-[propylcarbamoyl]oxirane-2-carbonyl)-*L*-isoleucyl-*L*-proline), CA-074-Me ((*L*-3-*trans*-[propylcarbamoyl]oxirane-2-carbonyl)-*L*-isoleucyl-*L*-proline-methylester), calpastatin, calpeptin (benzyloxycarbonylleucyl-norleucinal), carboxypeptidase inhibitor, cathepsin inhibitor I (Z-Phe-Gly-NHO-Bz), cathepsin inhibitor II (Z-Phe-Gly-NHO-Bz-*p*Me), cathepsin inhibitor III (Z-Phe-Gly-NHO-Bz-*p*OMe), cathepsin B inhibitor I (Z-Phe-Ala-CH<sub>2</sub>F), cathepsin B inhibitor II (Ac-Leu-Val-lysinal), cathepsin L inhibitor I (Z-Phe-Phe-CH<sub>2</sub>F), cathepsin L inhibitor II (Z-Phe-Tyr-CHO), cathepsin L inhibitor III (Z-Phe-Tyr-(*t*-Bu)-CHN<sub>2</sub>), cathepsin L inhibitor IV (1-naphthalenesulfonyl-Ile-Trp-CHO), cathepsin L inhibitor V (Z-Phe-Tyr(*O**t*Bu)-COCHO), cathepsin L inhibitor VI (*N*-(4-biphenylacetyl)-*S*-methyleysteine (*D*)-Arg-Phe-□-phenethylamide)(*N*-(4-biphenylacetyl)-*S*-methylcysteine-*(D*)-Arg-Phe- $\beta$ -phenethylamide), cathepsin S inhibitor (Z-Phe-Leu-COCHO), cystatin, diprotin A (H-Ile-Pro-Ile-OH), E-64 (*trans*-epoxysuccinyl-*L*-leucylamido-(4-guanidino)butane), E-64 d (loxistatin, or (2*S*,3*S*)-*trans*-epoxysuccinyl-*L*-leucylamido-3-methylbutane ethyl ester), ebelactone A (3,11-dihydroxy-2,4,6,8,10,12-hexamethyl-9-oxo-6-tetradecenoic 1,3-lactone), ebelactone B (2-ethyl-3,11-dihydroxy-4,6,8,10,12-penta methyl -9-oxo-6-~~

tetradecenoic 1,3-lactone), EDTA (ethylenediamine tetraacetic acid), EGTA (ethyleneglycol-bis(β-aminoethyl)-N,N,N',N'-tetraacetic acid) (ethyleneglycol-bis(β-aminoethyl)-N,N,N',N'-tetraacetic acid), elastase inhibitor II (MeOSuc-Ala-Ala-Pro-Ala-CMK), elastase inhibitor III (MeOSuc-Ala-Ala-Pro-Val-CMK), elastatinal (Leu-(Cap)-Gln-Ala-al or *N*-[(*S*)-1-carboxy-isopentyl]-carbamoyl-alpha-(2-iminohexahydro-4(*S*)-pyrimidyl]-*L*-glycyl-*L*-glutaminyl-*L*-alaninal), E-64 (*trans*-epoxysuccinyl-*L*-leucylamido-(4-guanidino)butane), E-64 d (loxistatin, or (2*S*,3*S*)-*trans*-epoxysuccinyl-*L*-leucylamido-3-methylbutane ethyl ester), *N*-ethyl maleimide, GGACK (1,5-dansyl-*L*-glutamyl-*L*-glycyl-*L*-arginine chloro methyl ketone), galardin (*N*-[(2*S*)-(methoxycarbonylmethyl)-4-methylpentanoyl]-*L*-tryptophan-methyl amide), 2-guanidinoethylmercaptosuccinic acid, hirudin, HIV protease inhibitor (Ac-Leu-Val-phenylalaninal), leuhistin (((2*R*,3*S*)-3-amino-2-hydroxy-2-(1*H*-imidazol-4-ylmethyl)-5-methyl)-5-methylhexanoic acid), leupeptin (acetyl-leucyl-leucyl-arginal), NCO-700, PEFABLOC SC (4-(2-aminoethyl)-benzenesulfonyl fluoride), pepstatin (isovaleryl-Val-Val-4-amino-3-hydroxy-6-methylheptanoyl-Ala-4-amino-3-hydroxy-6-methylheptanoic acid), phebestin ((2*S*,3*R*)-3-amino-2-hydroxy-4-phenylbutanoyl-*L*-valyl-*L*-phenylalanine), PMSF (phenyl methyl sulfonyl fluoride), phosphoramidon (*N*-alpha-*L*-rhamnopyranosyloxy(hydroxyl phosphinyl)-*L*-Leucyl-*L*-tryptophan, plummer's inhibitor (*D,L*-2-mercaptomethyl-3-guanidino-ethylthiopropanoic acid), 1,10-phenanthroline, subtilisin inhibitor I (Boc-Ala-Ala-NHO-Bz), subtilisin inhibitor II (Z-Gly-Phe-NHO-Bz), subtilisin inhibitor III (Z-Gly-Phe-NHO-Bz-*p*OMe), subtilisin inhibitor IV (Boc-Pro-Phe-NHO-Bz-*p*Cl), subtilisin inhibitor V (Boc-Ala-Pro-Phe-NHO-Bz), TIMP-2 (tissue inhibitor of metalloproteinase 2), trypsin inhibitor, secretory leukocyte protease inhibitor, and any mixture thereof.

20. (Original): The method of claim 18, wherein the agent that alters activities of G protein coupled receptors and cAMP or pharmaceutically acceptable derivative is selected from the group consisting of AB-MECA (*N*<sup>6</sup>-4-aminobenzyl-5'-*N*-methylcarboxamidoadenosine), CPA (*N*<sup>6</sup>-cyclopentyladenosine), ADAC (*N*<sup>6</sup>-[4-[[[4-[[[(2-aminoethyl) amino] carbonyl] methyl]-anilino] carbonyl] methyl] phenyl]adenosine), CCPA (2-chloro-*N*<sup>6</sup>-cyclopentyl adenosine), CHA (*N*<sup>6</sup>-cyclohexyladenosine),

GR79236 (*N*<sup>6</sup>-[1*S*, *trans*,2-hydroxycyclo pentyl] adenosine), *S*-ENBA ((2*S*)- *N*<sup>6</sup>-(2-endonorbanyl)adenosine), IAB-MECA (*N*<sup>6</sup>-(4-amino-3-iodobenzyl)adenosine-5'-*N*-methylcarboxamidoadenosine), *R*-PIA (*R*-*N*<sup>6</sup>-(phenyl isopropyl) adenosine), ATL146e (4-{3-[6-amino-9-(5-ethylcarbamoyl-3,4-dihydroxy-tetrahydro-furan-2-yl)-9H-purin-2-yl]-prop-2-ynyl}-cyclohexanecarboxylic acid methyl ester), CGS-21680 (APEC or 2-[*p*-(2-carbonyl-ethyl)-phenyl ethyl amino]-5'-*N*-ethylcarboxamidoadenosine), CV1808 (2-phenylaminoadenosine), HENECA (2-hex-1-ynyl-5'-*N*-ethylcarboxamido adenosine), NECA (5'-*N*-ethyl-carboxamido adenosine), PAPA-APEC (2-(4-[2-[(4-aminophenyl)methylcarbonyl]ethyl] phenyl) ethylamino-5'-*N*-ethyl carboxamidoadenosine), DITC-APEC (2-[*p*-(4-isothiocyanatophenylamino thio carbonyl-2-ethyl)-phenylethylamino]-5'-*N*-ethylcarboxamidoadenosine), DPMA (*N*<sup>6</sup>-(2(3,5-dimethoxy phenyl)-2-(2-methylphenyl)ethyl)adenosine), *S*-PHPNECA ((*S*)-2-phenylhydroxypropynyl-5'-*N*-ethylcarboxamidoadenosine), WRC-0470 (2-cyclohexyl methylidenehydrazinoadenosine), AMP-579 (1*S*-[1a,2b,3b,4a(*S*\*)])-4-[7-[[2-(3-chloro-2-thienyl)-1-methylpropyl]amino]-3H-imidazo [4,5-b] pyridyl-3-yl] cyclopentane carboxamide), IB-MECA (*N*<sup>6</sup>- (3-iodobenzyl)adenosine-5'-*N*-methyluronamide), 2-Cl-AADO (2-chloroadenosine), I-ABA (*N*<sup>6</sup>-(4-amino-3-iodobenzyl) adenosine), *S*-PIA (*S*-*N*<sup>6</sup>-(phenylisopropyl)adenosine), 2-[(2-aminoethyl-aminocarbonylethyl) phenylethyl amino]-5'-*N*-ethyl-carboxamidoadenosine, 2-Cl-IB-MECA (2-chloro-*N*<sup>6</sup>- (3-iodobenzyl) adenosine-5'-*N*-methyluronamide), adenosine, polyadenylic acid, and any mixture thereof.

21. (Original): A pharmaceutical composition comprising:
  - a. a protease inhibitor; and
  - b. an agent that alters activities of G protein coupled receptors and cAMP or a pharmaceutically acceptable derivative or prodrug thereof.
22. (Currently amended): The pharmaceutical composition of claim 21, wherein the protease inhibitor is selected from the group consisting of 4-(2-aminoethyl) benzenesulfonylfluoride, ~~amino-n-caproic acid-amino-n-caproic acid~~, ~~4-~~

antiehymotrypsin<sub>1</sub>-antichymotrypsin, antipain, antithrombin III,  $\square_1$ -antitrypsin<sub>1</sub>-antitrypsin, *p*-amidinophenylmethyl sulfonyl fluoride, aprotinin, cathepsin/subtilisin inhibitor (Boc-Val-Phe-NHO-Bz-*p*Cl), chymostatin ( $\{[(S)-1\text{-carboxy-2-phenylethyl}]\text{-carbamoyl-}\square-[2\text{-amidohexahydro-4(S)-pyrimidyl}]\text{-}(S)\text{-glycyl-[A=Leu, B=Val, or C=Ile]-phenylalaninal}\}[(S)-1\text{-carboxy-2-phenylethyl}]\text{-carbamoyl-}\alpha\text{-[2-amidohexahydro-4(S)-pyrimidyl]-}(S)\text{-glycyl-[A=Leu, B=Val, or C=Ile]-phenylalaninal}$ ), chymotrypsin inhibitor I, 3,4-dichloroisocoumarin, diisopropylfluoro phosphate, dipeptidylpeptidase IV inhibitor I (Ile-Pro-Ile), dipeptidylpeptidase IV inhibitor II (H-Glu-(NHO-Bz)-Pyr), ecotin, elastase inhibitor I (Boc-Ala-Ala-Ala-NHO-Bz),  $\square_2$ -macroglobulin<sub>2</sub>-macroglobulin, PPACK (*D*-Phe-Pro-Arg-chloromethylketone), PPACK II, *N*<sup>a</sup>-tosyl-Lys chloromethyl ketone, *N*<sup>a</sup>-tosyl-Phe chloromethyl ketone, acetyl-pepstatin (Ac-Val-Val-(3*S*,4*S*)-Sta-Ala-(3*S*,4*S*)-Sta-OH), calpain inhibitor I (*N*-acetyl-Leu-Leu-norleucinal), calpain inhibitor II (*N*-acetyl -Leu-Leu-Met-CHO), amastatin ( $[(2S, 2R)]\text{-3-amino-2-hydroxy-5-methylhexanoyl}-\text{Val-Val-Asp-OH}$ ), arphamenine A ((2*R*,5*S*)-5-amino-8-guanidino-4-oxo-2-phenylmethyl octanoic acid), arphamenine B ((2*R*,5*S*)-5-amino-8-guanidino-4-oxo-2-*p*-hydroxyphenyl methyloctanoic acid), benzamidine, bestatin ( $[(2S, 2R)]\text{-3-amino-2-hydroxy-4-phenyl butanoyl}-L\text{-Leucine}$ ), CA-074 ((*L*-3-*trans*-[propylcarbamoyl]oxirane-2-carbonyl)-*L*-isoleucyl-*L*-proline), CA-074-Me ((*L*-3-*trans*-[propylcarbamoyl]oxirane-2-carbonyl)-*L*-isoleucyl-*L*-proline-methylester), calpastatin, calpeptin (benzyloxycarbonylleucyl-norleucinal), carboxypeptidase inhibitor, cathepsin inhibitor I (Z-Phe-Gly-NHO-Bz), cathepsin inhibitor II (Z-Phe-Gly-NHO-Bz-*p*Me), cathepsin inhibitor III (Z-Phe-Gly-NHO-Bz-*p*OMe), cathepsin B inhibitor I (Z-Phe-Ala-CH<sub>2</sub>F), cathepsin B inhibitor II (Ac-Leu-Val-lysinal), cathepsin L inhibitor I (Z-Phe-Phe-CH<sub>2</sub>F), cathepsin L inhibitor II (Z-Phe-Tyr-CHO), cathepsin L inhibitor III (Z-Phe-Tyr-(*t*-Bu)-CHN<sub>2</sub>), cathepsin L inhibitor IV (1-naphthalenesulfonyl-Ile-Trp-CHO), cathepsin L inhibitor V (Z-Phe-Tyr(*t*Bu)-COCHO), cathepsin L inhibitor VI ( $N\text{-}(4\text{-biphenylacetyl})-S\text{-methyleysteine-(D)-Arg-Phe-}\square\text{-phenethylamide}(N\text{-}(4\text{-biphenylacetyl})-S\text{-methylcysteine-(D)-Arg-Phe-}\beta\text{-phenethylamide)}$ ), cathepsin S inhibitor (Z-Phe-Leu-COCHO), cystatin, diprotin A (H-Ile-Pro-Ile-OH), E-64 (*trans*-epoxysuccinyl-*L*-leucylamido-(4-guanidino)butane), E-64 d (loxistatin, or (2*S*,3*S*)-*trans*-epoxysuccinyl-*L*-

leucylamido-3-methylbutane ethyl ester), ebelactone A (3,11-dihydroxy-2,4,6,8,10,12-hexamethyl-9-oxo-6-tetradecenoic 1,3-lactone), ebelactone B (2-ethyl-3,11-dihydroxy-4,6,8,10,12-penta methyl -9-oxo-6-tetradecenoic 1,3-lactone), EDTA (ethylenediamine tetraacetic acid), EGTA (ethyleneglycol bis(β-aminoethyl)-N,N,N',N'-tetraacetic acid) (ethyleneglycol-bis(β-aminoethyl)-N,N,N',N'-tetraacetic acid), elastase inhibitor II (MeOSuc-Ala-Ala-Pro-Ala-CMK), elastase inhibitor III (MeOSuc-Ala-Ala-Pro-Val-CMK), elastatinal (Leu-(Cap)-Gln-Ala-al or *N*[(*S*)-1-carboxy-isopentyl]-carbamoyl-alpha-(2-iminohexahydro-4(*S*)-pyrimidyl]-*L*-glycyl-*L*-glutaminyl-*L*-alaninal), E-64 (*trans*-epoxysuccinyl-*L*-leucylamido-(4-guanidino)butane), E-64d (loxistatin, or (2*S*,3*S*)-*trans*-epoxysuccinyl-*L*-leucylamido-3-methylbutane ethyl ester), *N*-ethyl maleimide, GGACK (1,5-dansyl-*L*-glutamyl-*L*-glycyl-*L*-arginine chloro methyl ketone), galardin (*N*[(2*S*)-(methoxycarbonylmethyl)-4-methylpentanoyl]-*L*-tryptophan-methyl amide), 2-guanidinoethylmercaptosuccinic acid, hirudin, HIV protease inhibitor (Ac-Leu-Val-phenylalaninal), leuhistin (((2*R*,3*S*)-3-amino-2-hydroxy-2-(1*H*-imidazol-4-ylmethyl)-5-methyl)-5-methylhexanoic acid), leupeptin (acetyl-leucyl-leucyl-arginal), NCO-700, PEFABLOC SC (4-(2-aminoethyl)-benzenesulfonyl fluoride), pepstatin (isovaleryl-Val-Val-4-amino-3-hydroxy-6-methylheptanoyl-Ala-4-amino-3-hydroxy-6-methylheptanoic acid), phebestin ((2*S*,3*R*)-3-amino-2-hydroxy-4-phenylbutanoyl-*L*-valyl-*L*-phenylalanine), PMSF (phenyl methyl sulfonyl fluoride), phosphoramidon (*N*-alpha-*L*-rhamnopyranosyloxy(hydroxyl phosphinyl)-*L*-Leucyl-*L*-tryptophan, plummer's inhibitor (*D,L*-2-mercaptopethyl-3-guanidino-ethylthiopropanoic acid), 1,10-phenanthroline, subtilisin inhibitor I (Boc-Ala-Ala-NHO-Bz), subtilisin inhibitor II (Z-Gly-Phe-NHO-Bz), subtilisin inhibitor III (Z-Gly-Phe-NHO-Bz-*p*OMe), subtilisin inhibitor IV (Boc-Pro-Phe-NHO-Bz-*p*Cl), subtilisin inhibitor V (Boc-Ala-Pro-Phe-NHO-Bz), TIMP-2 (tissue inhibitor of metalloproteinase 2), trypsin inhibitor, secretory leukocyte protease inhibitor, and any mixture there of.

23. (Original): The pharmaceutical composition of claim 21, wherein the agent that alters activities of G protein coupled receptors and cAMP or pharmaceutically acceptable derivative is selected from the group consisting of AB-MECA (*N*<sup>6</sup>-4-aminobenzyl-5'-*N*-

methylcarbox amidoadenosine), CPA ( $N^6$ -cyclopentyladenosine), ADAC ( $N^6$ -[4-[[[4-  
[[[(2-aminoethyl) amino] carbonyl] methyl]-anilino] carbonyl] methyl] phenyl]  
adenosine), CCPA (2-chloro- $N^6$ -cyclopentyladenosine), CHA ( $N^6$ -cyclohexyladenosine),  
GR79236 ( $N^6$ -[1*S*, *trans*,2-hydroxycyclopentyl] adenosine), *S*-ENBA ((2*S*)-  $N^6$ -(2-  
endonorbanyl)adenosine), IAB-MECA ( $N^6$ -(4-amino-3-iodobenzyl)adenosine-5'-*N*-  
methylcarboxamidoadenosine), *R*-PIA (*R*- $N^6$ -(phenylisopropyl) adenosine), ATL146e (4-  
{3-[6-amino-9-(5-ethyl carbamoyl -3,4-dihydroxy-tetrahydro-furan-2-yl)-9H-purin-2-yl]-  
prop-2-ynyl}-cyclohexanecarboxylic acid methyl ester), CGS-21680 (APEC or 2-[*p*-(2-  
carbonyl-ethyl)-phenyl ethyl amino]-5'-*N*-ethylcarboxamidoadenosine), CV1808 (2-  
phenylamino adenosine, HENECA (2-hex-1-ynyl-5'-*N*-ethylcarboxamido adenosine),  
NECA (5'-*N*-ethyl-carboxamido adenosine), PAPA-APEC (2-(4-[2-[(4-  
aminophenyl)methylcarbonyl] ethyl] phenyl ethylamino-5'-*N*-ethyl  
carboxamidoadenosine), DITC-APEC (2-[*p*-(4-isothiocyanatophenylamino thiocabonyl-  
2-ethyl)-phenylethylamino]-5'-*N*-ethylcarbox amido adenosine), DPMA ( $N^6$ -(2(3,5-  
dimethoxy phenyl)-2-(2-methylphenyl)ethyl) adenosine), *S*-PHPNECA ((*S*)-2-  
phenylhydroxypropynyl-5'-*N*-ethylcarboxamido adenosine), WRC-0470 (2-  
cyclohexylmethylenedihydrazinoadenosine), AMP-579 (1*S*-[1a,2b,3b,4a(S\*)])-4-[7-[[2-  
(3-chloro-2-thienyl)-1-methylpropyl]amino]-3H-imidazo [4,5-b] pyridyl-3-yl]  
cyclopentane carboxamide), IB-MECA ( $N^6$ - (3-iodobenzyl) adenosine -5'-*N*-  
methyluronamide), 2-CIADO (2-chloroadenosine), I-ABA ( $N^6$ -(4-amino-3-iodobenzyl)  
adenosine), *S*-PIA (*S*- $N^6$ -(phenylisopropyl)adenosine), 2-[(2-aminoethyl-  
aminocarbonylethyl) phenylethyl amino]-5'-*N*-ethyl-carboxamidoadenosine, 2-Cl-IB-  
MECA (2-chloro- $N^6$ - (3-iodobenzyl)adenosine-5'-*N*-methyluronamide), adenosine,  
polyadenylic acid, and any mixture thereof.

24. (Original): A method of preventing organ ischemia or reperfusion injury comprising  
concomitantly administering to a living subject in need thereof

- a. a protease inhibitor; and
- b. an agent that alters activities of G protein coupled receptors and cAMP or a  
pharmaceutically acceptable derivative or prodrug thereof.

25. (Currently amended): The method of claim 24, wherein the protease inhibitor is selected from the group consisting of 4-(2-aminoethyl) benzenesulfonylfluoride, ~~□-amino-n-caproic acid, □-antichymotrypsin~~<sub>1</sub>-~~antichymotrypsin~~, antipain, antithrombin III, ~~□-antitrypsin~~<sub>1</sub>-~~antitrypsin~~, *p*-amidinophenylmethyl sulfonyl fluoride, aprotinin, cathepsin/subtilisin inhibitor (Boc-Val-Phe-NHO-Bz-*p*Cl), chymostatin (~~[(S)-1-carboxy-2-phenylethyl]-carbamoyl-□-[2-amidohexahydro-4(S)-pyrimidyl]-~~(*S*)-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal)~~[(S)-1-carboxy-2-phenylethyl]-carbamoyl-α-[2-amidohexahydro-4(S)-pyrimidyl]-~~(*S*)-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal), chymotrypsin inhibitor I, 3,4-dichloroisocoumarin, diisopropylfluoro phosphate, dipeptidylpeptidase IV inhibitor I (Ile-Pro-Ile), dipeptidylpeptidase IV inhibitor II (H-Glu-(NHO-Bz)-Pyr), ecatin, elastase inhibitor I (Boc-Ala-Ala-Ala-NHO-Bz), ~~□-macroglobulin~~<sub>2</sub>-~~macroglobulin~~, PPACK (*D*-Phe-Pro-Arg-chloromethylketone), PPACK II, *N*<sup>a</sup>-tosyl-Lys chloromethyl ketone, *N*<sup>a</sup>-tosyl-Phe chloromethyl ketone, acetyl-pepstatin (Ac-Val-Val-(3*S*,4*S*)-Sta-Ala-(3*S*,4*S*)-Sta-OH), calpain inhibitor I (*N*-acetyl-Leu-Leu-norleucinal), calpain inhibitor II (*N*-acetyl-Leu-Leu-Met-CHO), amastatin (~~[(2*S*, 2*R*)]-3-amino-2-hydroxy-5-methylhexanoyl~~-Val-Val-Asp-OH), arphamenine A ((2*R*,5*S*)-5-amino-8-guanidino-4-oxo-2-phenylmethyl octanoic acid), arphamenine B ((2*R*,5*S*)-5-amino-8-guanidino-4-oxo-2-*p*-hydroxyphenyl methyloctanoic acid), benzamidine, bestatin (~~[(2*S*, 2*R*)-3-amino-2-hydroxy-4-phenyl butanoyl]~~-*L*-Leucine), CA-074 ((*L*-3-*trans*-[propylcarbamoyl]oxirane-2-carbonyl)-*L*-isoleucyl-*L*-proline), CA-074-Me ((*L*-3-*trans*-[propylcarbamoyl]oxirane-2-carbonyl)-*L*-isoleucyl-*L*-proline-methylester), calpastatin, calpeptin (benzyloxycarbonylleucyl-norleucinal), carboxypeptidase inhibitor, cathepsin inhibitor I (Z-Phe-Gly-NHO-Bz), cathepsin inhibitor II (Z-Phe-Gly-NHO-Bz-*p*Me), cathepsin inhibitor III (Z-Phe-Gly-NHO-Bz-*p*OMe), cathepsin B inhibitor I (Z-Phe-Ala-CH<sub>2</sub>F), cathepsin B inhibitor II (Ac-Leu-Val-lysinal), cathepsin L inhibitor I (Z-Phe-Phe-CH<sub>2</sub>F), cathepsin L inhibitor II (Z-Phe-Tyr-CHO), cathepsin L inhibitor III (Z-Phe-Tyr-(*t*-Bu)-CHN<sub>2</sub>), cathepsin L inhibitor IV (1-naphthalenesulfonyl-Ile-Trp-CHO), cathepsin L inhibitor V (Z-Phe-Tyr(O/Bu)-COCHO), cathepsin L inhibitor VI (*N*-(4-biphenylacetyl)-*S*-methyleysteine (*D*)-Arg-Phe-□-

phenethylamide)(N-(4-biphenylacetyl)-S-methylcysteine-(D)-Arg-Phe-β-phenethylamide), cathepsin S inhibitor (Z-Phe-Leu-COCHO), cystatin, diprotin A (H-Ile-Pro-Ile-OH), E-64 (*trans*-epoxysuccinyl-*L*-leucylamido-(4-guanidino)butane), E-64 d (loxistatin, or (2*S*,3*S*)-*trans*-epoxysuccinyl-*L*-leucylamido-3-methylbutane ethyl ester), ebelactone A (3,11-dihydroxy-2,4,6,8,10,12-hexamethyl-9-oxo-6-tetradecenoic 1,3-lactone), ebelactone B (2-ethyl-3,11-dihydroxy-4,6,8,10,12-penta methyl -9-oxo-6-tetradecenoic 1,3-lactone), EDTA (ethylenediamine tetraacetic acid), EGTA (ethyleneglycol-bis(ε-aminoethyl)-N,N,N',N'-tetraacetic acid)(ethyleneglycol-bis(β-aminoethyl)-N,N,N',N'-tetraacetic acid), elastase inhibitor II (MeOSuc-Ala-Ala-Pro-Ala-CMK), elastase inhibitor III (MeOSuc-Ala-Ala-Pro-Val-CMK), elastatinal (Leu-(Cap)-Gln-Ala-al or *N*-[(*S*)-1-carboxy-isopentyl]-carbamoyl-alpha-(2-iminohexahydro-4(*S*)-pyrimidyl]-*L*-glycyl-*L*-glutaminyl-*L*-alaninal), E-64 (*trans*-epoxysuccinyl-*L*-leucylamido-(4-guanidino)butane), E-64d (loxistatin, or (2*S*,3*S*)-*trans*-epoxysuccinyl-*L*-leucylamido-3-methylbutane ethyl ester), *N*-ethyl maleimide, GGACK (1,5-dansyl-*L*-glutamyl-*L*-glycyl-*L*-arginine chloro methyl ketone), galardin (*N*-[(2*S*)-(methoxycarbonylmethyl)-4-methylpentanoyl]-*L*-tryptophan-methyl amide), 2-guanidinoethylmercaptosuccinic acid, hirudin, HIV protease inhibitor (Ac-Leu-Val-phenylalaninal), leuhistin (((2*R*,3*S*)-3-amino-2-hydroxy-2-(1*H*-imidazol-4-ylmethyl)-5-methyl)-5-methylhexanoic acid), leupeptin (acetyl-leucyl-leucyl-arginal), NCO-700, PEFALOC SC (4-(2-aminoethyl)-benzenesulfonyl fluoride), pepstatin (isovaleryl-Val-Val-4-amino-3-hydroxy-6-methylheptanoyl-Ala-4-amino-3-hydroxy-6-methylheptanoic acid), phebestin ((2*S*,3*R*)-3-amino-2-hydroxy-4-phenylbutanoyl-*L*-valyl-*L*-phenylalanine), PMSF (phenyl methyl sulfonyl fluoride), phosphoramidon (*N*-alpha-*L*-rhamnopyranosyloxy(hydroxyl phosphinyl)-*L*-Leucyl-*L*-tryptophan, plummer's inhibitor (*D,L*-2-mercaptopethyl-3-guanidino-ethylthiopropanoic acid), 1,10-phenanthroline, subtilisin inhibitor I (Boc-Ala-Ala-NHO-Bz), subtilisin inhibitor II (Z-Gly-Phe-NHO-Bz), subtilisin inhibitor III (Z-Gly-Phe-NHO-Bz-*p*OMe), subtilisin inhibitor IV (Boc-Pro-Phe-NHO-Bz-*p*Cl), subtilisin inhibitor V (Boc-Ala-Pro-Phe-NHO-Bz), TIMP-2 (tissue inhibitor of metalloproteinase 2), trypsin inhibitor, secretory leukocyte protease inhibitor, and any mixture thereof.

26. (Original): The method of claim 24, wherein the agent that alters the activities of G-protein coupled receptors and cAMP or pharmaceutically acceptable derivative is selected from the group consisting of AB-MECA ( $N^6$ -4-aminobenzyl-5'- $N$ -methylcarboxamidoadenosine), CPA ( $N^6$ -cyclopentyladenosine), ADAC ( $N^6$ -[4-[[[4-[(2-aminoethyl) amino] carbonyl] methyl]-anilino] carbonyl] methyl] phenyl] adenosine), CCPA (2-chloro- $N^6$ -cyclopentyl adenosine), CHA ( $N^6$ -cyclohexyladenosine), GR79236 ( $N^6$ -[1*S*, *trans*,2-hydroxycyclo pentyl] adenosine), S-ENBA ((2*S*)-  $N^6$ -(2-endonorbanyl)adenosine), IAB-MECA ( $N^6$ -(4-amino-3-iodobenzyl)adenosine-5'- $N$ -methylcarboxamidoadenosine), *R*-PIA (*R*- $N^6$ -(phenyl isopropyl) adenosine), ATL146e (4-{3-[6-amino-9-(5-ethylcarbamoyl-3,4-dihydroxy-tetrahydro-furan-2-yl)-9H-purin-2-yl]-prop-2-ynyl}-cyclohexanecarboxylic acid methyl ester), CGS-21680 (APEC or 2-[*p*-(2-carbonyl-ethyl)-phenyl ethyl amino]-5'- $N$ -ethylcarboxamidoadenosine), CV1808 (2-phenylaminoadenosine), HENECA (2-hex-1-ynyl-5'- $N$ -ethylcarboxamido adenosine), NECA (5'- $N$ -ethyl-carboxamido adenosine), PAPA-APEC (2-(4-[2-[(4-aminophenyl)methylcarbonyl]ethyl] phenyl) ethylamino-5'- $N$ -ethyl carboxamidoadenosine), DITC-APEC (2-[*p*-(4-isothiocyanatophenylamino thio carbonyl-2-ethyl)-phenylethylamino]-5'- $N$ -ethylcarboxamidoadenosine), DPMA ( $N^6$ -(2(3,5-dimethoxy phenyl)-2-(2-methylphenyl)ethyl)adenosine), S-PHPNECA ((*S*)-2-phenylhydroxypropynyl-5'- $N$ -ethylcarboxamidoadenosine), WRC-0470 (2-cyclohexyl methylidenehydrazinoadenosine), AMP-579 (1*S*-[1a,2b,3b,4a(*S*\*)])-4-[7-[[2-(3-chloro-2-thienyl)-1-methylpropyl]amino]-3H-imidazo [4,5-b] pyridyl-3-yl] cyclopentane carbox amide), IB-MECA ( $N^6$ -(3-iodobenzyl)adenosine-5'- $N$ -methyluronamide), 2-CIADO (2-chloroadenosine), I-ABA ( $N^6$ -(4-amino-3-iodobenzyl) adenosine), *S*-PIA (*S*- $N^6$ -(phenyl isopropyl) adenosine), 2-[(2-aminoethyl-aminocarbonylethyl) phenylethyl amino]-5'- $N$ -ethyl-carboxamidoadenosine, 2-Cl-IB-MECA (2-chloro- $N^6$ -(3-iodobenzyl)adenosine-5'- $N$ -methyluronamide), adenosine, polyadenylic acid, and any mixture thereof.

27. (Original): A method of preventing organ ischemia or reperfusion injury comprising administering to a living subject in need thereof sequentially in any order

a. a protease inhibitor; and

b. an agent that alters activities of G protein coupled receptors and cAMP or a pharmaceutically acceptable derivative or prodrug thereof.

28. (Currently amended): The method of claim 27, wherein the serine protease inhibitor is selected from the group consisting of 4-(2-aminoethyl) benzenesulfonylfluoride,  $\alpha$ -amino-n-caproic acid-amino-n-caproic acid,  $\alpha$ -antichymotrypsin $\alpha$ <sub>1</sub>-antichymotrypsin, antipain, antithrombin III,  $\alpha$ <sub>1</sub>-antitrypsin $\alpha$ <sub>1</sub>-antitrypsin, *p*-amidinophenylmethyl sulfonyl fluoride, aprotinin, cathepsin/subtilisin inhibitor (Boc-Val-Phe-NHO-Bz-*p*Cl), chymostatin (~~[(S)-1-carboxy-2-phenylethyl]-carbamoyl-α-[2-amidohexahydro-4(S)-pyrimidyl]-*S*-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal~~)~~[(S)-1-carboxy-2-phenylethyl]-carbamoyl-α-[2-amidohexahydro-4(S)-pyrimidyl]-*S*-glycyl-[A = Leu, B = Val, or C = Ile]-phenylalaninal~~, chymotrypsin inhibitor I, 3,4-dichloroisocoumarin, diisopropylfluoro phosphate, dipeptidylpeptidase IV inhibitor I (Ile-Pro-Ile), dipeptidylpeptidase IV inhibitor II (H-Glu-(NHO-Bz)-Pyr), ecotin, elastase inhibitor I (Boc-Ala-Ala-Ala-NHO-Bz),  $\alpha$ <sub>2</sub>-macroglobulin $\alpha$ <sub>2</sub>-macroglobulin, PPACK (*D*-Phe-Pro-Arg-chloromethylketone), PPACK II, *N*<sup>a</sup>-tosyl-Lys chloromethyl ketone, *N*<sup>a</sup>-tosyl-Phe chloromethyl ketone, acetyl-pepstatin (Ac-Val-Val-(3*S*,4*S*)-Sta-Ala-(3*S*,4*S*)-Sta-OH), calpain inhibitor I (*N*-acetyl-Leu-Leu-norleucinal), calpain inhibitor II (*N*-acetyl-Leu-Leu-Met-CHO), amastatin (~~[(2*S*,2*R*)]-3-amino-2-hydroxy-5-methylhexanoyl~~-Val-Val-Asp-OH), arphamenine A ((2*R*,5*S*)-5-amino-8-guanidino-4-oxo-2-phenylmethyl octanoic acid), arphamenine B ((2*R*,5*S*)-5-amino-8-guanidino-4-oxo-2-*p*-hydroxyphenyl methyloctanoic acid), benzamidine, bestatin (~~[(2*S*,2*R*)-3-amino-2-hydroxy-4-phenyl butanoyl]-*L*-Leucine~~), CA-074 ((*L*-3-*trans*-[propylcarbamoyl]oxirane-2-carbonyl)-*L*-isoleucyl-*L*-proline), CA-074-Me ((*L*-3-*trans*-[propylcarbamoyl]oxirane-2-carbonyl)-*L*-isoleucyl-*L*-proline-methylester), calpastatin, calpeptin (benzyloxycarbonylleucyl-norleucinal), carboxypeptidase inhibitor, cathepsin inhibitor I (Z-Phe-Gly-NHO-Bz), cathepsin inhibitor II (Z-Phe-Gly-NHO-Bz-*p*Me), cathepsin inhibitor III (Z-Phe-Gly-NHO-Bz-*p*OMe), cathepsin B inhibitor I (Z-Phe-Ala-CH<sub>2</sub>F), cathepsin B inhibitor II (Ac-Leu-Val-lysinal), cathepsin L inhibitor I (Z-Phe-Phe-CH<sub>2</sub>F), cathepsin L inhibitor II (Z-Phe-Tyr-CHO), cathepsin L inhibitor III (Z-Phe-Tyr-(*t*-Bu)-CHN<sub>2</sub>), cathepsin L inhibitor

IV (1-naphthalenesulfonyl-Ile-Trp-CHO), cathepsin L inhibitor V (Z-Phe-Tyr(OtBu)-COCHO), cathepsin L inhibitor VI (*N*-(4-biphenylacetyl)-*S*-methylcysteine-(*D*)-Arg-Phe- $\square$ -phenethylamide)(*N*-(4-biphenylacetyl)-*S*-methylcysteine-(*D*)-Arg-Phe- $\beta$ -phenethylamide), cathepsin S inhibitor (Z-Phe-Leu-COCHO), cystatin, diprotin A (H-Ile-Pro-Ile-OH), E-64 (*trans*-epoxysuccinyl-*L*-leucylamido-(4-guanidino)butane), E-64 d (loxistatin, or (2*S*,3*S*)-*trans*-epoxysuccinyl-*L*-leucylamido-3-methylbutane ethyl ester), ebelactone A (3,11-dihydroxy-2,4,6,8,10,12-hexamethyl-9-oxo-6-tetradecenoic 1,3-lactone), ebelactone B (2-ethyl-3,11-dihydroxy-4,6,8,10,12-penta methyl -9-oxo-6-tetradecenoic 1,3-lactone), EDTA (ethylenediamine tetraacetic acid), EGTA (*ethyleneglycol-bis*( $\square$ -aminoethyl)-*N,N,N',N'*-tetraacetic acid)(*ethyleneglycol-bis*( $\beta$ -aminoethyl)-*N,N,N',N'*-tetraacetic acid), elastase inhibitor II (MeOSuc-Ala-Ala-Pro-Ala-CMK), elastase inhibitor III (MeOSuc-Ala-Ala-Pro-Val-CMK), elastatinal (Leu-(Cap)-Gln-Ala-al or *N*-[(*S*)-1-carboxy-isopentyl]-carbamoyl-alpha-(2-iminohexahydro-4(*S*)-pyrimidyl]-*L*-glycyl-*L*-glutaminyl-*L*-alaninal), E-64 (*trans*-epoxysuccinyl-*L*-leucylamido-(4-guanidino)butane), E-64d (loxistatin, or (2*S*,3*S*)-*trans*-epoxysuccinyl-*L*-leucylamido-3-methylbutane ethyl ester), *N*-ethyl maleimide, GGACK (1,5-dansyl-*L*-glutamyl-*L*-glycyl-*L*-arginine chloro methyl ketone), galardin (*N*-[(2*S*)-(methoxycarbonylmethyl)-4-methylpentanoyl]-*L*-tryptophan-methyl amide), 2-guanidinoethylmercaptosuccinic acid, hirudin, HIV protease inhibitor (Ac-Leu-Val-phenylalaninal), leuhistin (((2*R*,3*S*)-3-amino-2-hydroxy-2-(1*H*-imidazol-4-ylmethyl)-5-methyl)-5-methylhexanoic acid), leupeptin (acetyl-leucyl-leucyl-arginal), NCO-700, PEFABLOC SC (4-(2-aminoethyl)-benzenesulfonyl fluoride), pepstatin (isovaleryl-Val-Val-4-amino-3-hydroxy-6-methylheptanoyl-Ala-4-amino-3-hydroxy-6-methylheptanoic acid), phebestin ((2*S*,3*R*)-3-amino-2-hydroxy-4-phenylbutanoyl-*L*-valyl-*L*-phenylalanine), PMSF (phenyl methyl sulfonyl fluoride), phosphoramidon (*N*-alpha-*L*-rhamnopyranosyloxy(hydroxyl phosphinyl)-*L*-Leucyl-*L*-tryptophan, plummer's inhibitor (*D,L*-2-mercaptomethyl-3-guanidino-ethylthiopropanoic acid), 1,10-phenanthroline, subtilisin inhibitor I (Boc-Ala-Ala-NHO-Bz), subtilisin inhibitor II (Z-Gly-Phe-NHO-Bz), subtilisin inhibitor III (Z-Gly-Phe-NHO-Bz-*p*OMe), subtilisin inhibitor IV (Boc-Pro-Phe-NHO-Bz-*p*Cl), subtilisin inhibitor V (Boc-Ala-Pro-Phe-NHO-Bz), TIMP-2 (tissue inhibitor of metalloproteinase

2), trypsin inhibitor, secretory leukocyte protease inhibitor, and any mixture thereof.

29. (Original): The method of claim 27, wherein the agent that alters activities of G-protein coupled receptors and cAMP or pharmaceutically acceptable derivative is selected from the group consisting of AB-MECA ( $N^6$ -4-aminobenzyl-5'- $N$ -methylcarboxamidoadenosine), CPA ( $N^6$ -cyclopentyladenosine), ADAC ( $N^6$ -[4-[[[4-[(2-aminoethyl) amino] carbonyl] methyl]-anilino] carbonyl] methyl] phenyl] adenosine), CCPA (2-chloro- $N^6$ -cyclopentyl adenosine), CHA ( $N^6$ -cyclohexyladenosine), GR79236 ( $N^6$ -[1*S*, *trans*,2-hydroxycyclo pentyl] adenosine), S-ENBA ((2*S*)-  $N^6$ -(2-endonorbanyl)adenosine), IAB-MECA ( $N^6$ -(4-amino-3-iodobenzyl)adenosine-5'- $N$ -methylcarboxamidoadenosine), *R*-PIA (*R*- $N^6$ -(phenyl isopropyl) adenosine), ATL146e (4-{3-[6-amino-9-(5-ethylcarbamoyl-3,4-dihydroxy-tetrahydro-furan-2-yl)-9H-purin-2-yl]-prop-2-ynyl}-cyclohexanecarboxylic acid methyl ester), CGS-21680 (APEC or 2-[*p*-(2-carbonyl-ethyl)-phenyl ethyl amino]-5'- $N$ -ethylcarboxamidoadenosine), CV1808 (2-phenylaminoadenosine), HENECA (2-hex-1-ynyl-5'- $N$ -ethylcarboxamido adenosine), NECA (5'- $N$ -ethyl-carboxamido adenosine), PAPA-APEC (2-(4-[2-[(4-aminophenyl)methylcarbonyl]ethyl] phenyl) ethylamino-5'- $N$ -ethyl carboxamidoadenosine), DITC-APEC (2-[*p*-(4-isothiocyanatophenylamino thio carbonyl-2-ethyl)-phenylethylamino]-5'- $N$ -ethylcarboxamidoadenosine), DPMA ( $N^6$ -(2(3,5-dimethoxy phenyl)-2-(2-methylphenyl)ethyl)adenosine), *S*-PHPNECA ((*S*)-2-phenylhydroxypropynyl-5'- $N$ -ethylcarboxamidoadenosine), WRC-0470 (2-cyclohexyl methyldenehydrazinoadenosine), AMP-579 (1*S*-[1a,2b,3b,4a(S\*)])-4-[7-[[2-(3-chloro-2-thienyl)-1-methylpropyl]amino]-3H-imidazo [4,5-b] pyridyl-3-yl] cyclopentane carboxamide), IB-MECA ( $N^6$ - (3-iodobenzyl)adenosine-5'- $N$ -methyluronamide), 2-ClADO (2-chloroadenosine), I-ABA ( $N^6$ -(4-amino-3-iodobenzyl) adenosine), *S*-PIA (*S*- $N^6$ -(phenylisopropyl)adenosine), 2-[(2-aminoethyl-aminocarbonylethyl) phenylethyl amino]-5'- $N$ -ethyl-carboxamidoadenosine, 2-Cl-IB-MECA (2-chloro- $N^6$ - (3-iodobenzyl) adenosine-5'- $N$ -methyluronamide), adenosine, polyadenylic acid, and any mixture thereof.

30. (Original): A method of preventing organ or tissue injury at predetermined point or period of intervention comprising administrating to a living subject in need thereof a pharmaceutical composition comprising:
  - a. a protease inhibitor; and
  - b. an agent that alters activities of G protein coupled receptors and cAMP, an analog or a pharmaceutically acceptable derivative or prodrug thereof.
31. (Original): The method of claim 30, wherein the organ or tissue injury is related to at least one of cardiac surgery, non-surgical cardiac revascularization, organ transplantation, perfusion, ischemia, reperfusion, ischemia-reperfusion injury, oxidant injury, cytokine induced injury, shock induced injury, resuscitations injury, or apoptosis.
32. (Original): The method of claim 30, wherein the administration is made at the predetermined point of time related to at least one of pre-treatment regimen, pharmacological preconditioning, reperfusion or post interventional therapy, wherein the pharmacological preconditioning is a treatment administered before the ischemic intervention followed by a brief period of reperfusion or washout.
33. (Currently amended): The method of claim 30, wherein the protease inhibitor is selected from the group consisting of 4-(2-aminoethyl) benzenesulfonylfluoride, ~~□-amino-n-caproic acid~~, ~~□-antichymotrypsin~~ $\alpha_1$ -antichymotrypsin, antipain, antithrombin III, ~~□-antitrypsin~~ $\alpha_1$ -antitrypsin, *p*-amidinophenylmethyl sulfonyl fluoride, aprotinin, cathepsin/subtilisin inhibitor (Boc-Val-Phe-NHO-Bz-*p*Cl), chymostatin  $\{[(S)-1\text{-carboxy-2-phenylethyl}]\text{-carbamoyl-}[(2\text{-amidohexahydro-4}(S)\text{-pyrimidyl})\text{-}(S)\text{-glycyl-}[(A = \text{Leu, B = Val, or C = Ile})\text{-phenylalaninal}][(S)-1\text{-carboxy-2-phenylethyl}]\text{-carbamoyl-}[(A = \text{Leu, B = Val, or C = Ile})\text{-phenylalaninal}]\}$ , chymotrypsin inhibitor I, 3,4-dichloroisocoumarin, diisopropylfluoro phosphate, dipeptidylpeptidase IV inhibitor I (Ile-Pro-Ile), dipeptidylpeptidase IV inhibitor II (H-Glu-(NHO-Bz)-Pyr), ecotin, elastase inhibitor I (Boc-Ala-Ala-Ala-NHO-Bz), ~~□-2-macroglobulin~~ $\alpha_2$ -macroglobulin, PPACK (*D*-Phe-Pro-Arg-chloromethylketone),

PPACK II, *N*<sup>2</sup>-tosyl-Lys chloromethyl ketone, *N*<sup>2</sup>-tosyl-Phe chloromethyl ketone, acetyl-pepstatin (Ac-Val-Val-(3*S*,4*S*)-Sta-Ala-(3*S*,4*S*)-Sta-OH), calpain inhibitor I (*N*-acetyl-Leu-Leu-norleucinal), calpain inhibitor II (*N*-acetyl-Leu-Leu-Met-CHO), amastatin ([(2*S*, 2*R*)]-3-amino-2-hydroxy-5-methylhexanoyl] -Val-Val-Asp-OH), arphamenine A ((2*R*,5*S*)-5-amino-8-guanidino-4-oxo-2-phenylmethyl octanoic acid), arphamenine B ((2*R*,5*S*)-5-amino-8-guanidino-4-oxo-2-*p*-hydroxyphenyl methyloctanoic acid), benzamidine, bestatin ([(2*S*, 2*R*)-3-amino-2-hydroxy-4-phenyl butanoyl] -*L*-Leucine), CA-074 ((*L*-3-*trans*-[propylcarbamoyl]oxirane-2-carbonyl)-*L*-isoleucyl-*L*-proline), CA-074-Me ((*L*-3-*trans*-[propylcarbamoyl]oxirane-2-carbonyl)-*L*-isoleucyl-*L*-proline-methylester), calpastatin, calpeptin (benzyloxycarbonylleucyl-norleucinal), carboxypeptidase inhibitor, cathepsin inhibitor I (Z-Phe-Gly-NHO-Bz), cathepsin inhibitor II (Z-Phe-Gly-NHO-Bz-*p*Me), cathepsin inhibitor III (Z-Phe-Gly-NHO-Bz-*p*OMe), cathepsin B inhibitor I (Z-Phe-Ala-CH<sub>2</sub>F), cathepsin B inhibitor II (Ac-Leu-Val-lysinal), cathepsin L inhibitor I (Z-Phe-Phe-CH<sub>2</sub>F), cathepsin L inhibitor II (Z-Phe-Tyr-CHO), cathepsin L inhibitor III (Z-Phe-Tyr-(*t*-Bu)-CHN<sub>2</sub>), cathepsin L inhibitor IV (1-naphthalenesulfonyl-Ile-Trp-CHO), cathepsin L inhibitor V (Z-Phe-Tyr(O*t*Bu)-COCHO), cathepsin L inhibitor VI (*N*-(4-biphenylacetyl)-*S*-methyleysteine (*D*)-Arg-Phe- $\square$ -phenethylamide)(*N*-(4-biphenylacetyl)-*S*-methylcysteine-(*D*)-Arg-Phe- $\beta$ -phenethylamide), cathepsin S inhibitor (Z-Phe-Leu-COCHO), cystatin, diprotin A (H-Ile-Pro-Ile-OH), E-64 (*trans*-epoxysuccinyl-*L*-leucylamido-(4-guanidino)butane), E-64 d (loxistatin, or (2*S*,3*S*)-*trans*-epoxysuccinyl-*L*-leucylamido-3-methylbutane ethyl ester), ebelactone A (3,11-dihydroxy-2,4,6,8,10,12-hexamethyl-9-oxo-6-tetradecenoic 1,3-lactone), ebelactone B (2-ethyl-3,11-dihydroxy-4,6,8,10,12-penta methyl -9-oxo-6-tetradecenoic 1,3-lactone), EDTA (ethylenediamine tetraacetic acid), EGTA (ethyleneglycol-bis( $\square$ -aminoethyl)-*N,N,N',N'*-tetraacetic acid)(ethyleneglycol-bis( $\beta$ -aminoethyl)-*N,N,N',N'*-tetraacetic acid), elastase inhibitor II (MeOSuc-Ala-Ala-Pro-Ala-CMK), elastase inhibitor III (MeOSuc-Ala-Ala-Pro-Val-CMK), elastatinal (Leu-(Cap)-Gln-Ala-al or *N*-[(*S*)-1-carboxy-isopentyl]-carbamoyl-alpha-(2-iminohexahydro-4(*S*)-pyrimidyl]-*L*-glycyl-*L*-glutaminyl-*L*-alaninal), E-64 (*trans*-epoxysuccinyl-*L*-leucylamido-(4-guanidino)butane), E-64d (loxistatin, or (2*S*,3*S*)-*trans*-epoxysuccinyl-*L*-leucylamido-

3-methylbutane ethyl ester), *N*-ethyl maleimide, GGACK (1,5-dansyl-*L*-glutamyl-*L*-glycyl-*L*-arginine chloro methyl ketone), galardin (*N*-[(2*S*)-(methoxycarbonylmethyl)-4-methylpentanoyl]-*L*-tryptophan-methyl amide), 2-guanidinoethylmercaptosuccinic acid, hirudin, HIV protease inhibitor (Ac-Leu-Val-phenylalaninal), leuhistin (((2*R*,3*S*)-3-amino-2-hydroxy-2-(1*H*-imidazol-4-ylmethyl)-5-methyl)-5-methylhexanoic acid), leupeptin (acetyl-leucyl-leucyl-arginal), NCO-700, PEFALOC SC (4-(2-aminoethyl)-benzenesulfonyl fluoride), pepstatin (isovaleryl-Val-Val-4-amino-3-hydroxy-6-methylheptanoyl-Ala-4-amino-3-hydroxy-6-methylheptanoic acid), phebestin ((2*S*,3*R*)-3-amino-2-hydroxy-4-phenylbutanoyl-*L*-valyl-*L*-phenylalanine), PMSF (phenyl methyl sulfonyl fluoride), phosphoramidon (*N*-alpha-*L*-rhamnopyranosyloxy(hydroxyl phosphinyl)-*L*-Leucyl-*L*-tryptophan, plummer's inhibitor (*D,L*-2-mercaptomethyl-3-guanidino-ethylthiopropanoic acid), 1,10-phenanthroline, subtilisin inhibitor I (Boc-Ala-Ala-NHO-Bz), subtilisin inhibitor II (Z-Gly-Phe-NHO-Bz), subtilisin inhibitor III (Z-Gly-Phe-NHO-Bz-*p*OMe), subtilisin inhibitor IV (Boc-Pro-Phe-NHO-Bz-*p*Cl), subtilisin inhibitor V (Boc-Ala-Pro-Phe-NHO-Bz), TIMP-2 (tissue inhibitor of metalloproteinase 2), trypsin inhibitor, secretory leukocyte protease inhibitor, and any mixture thereof.

34. (Original): The method of claim 30, wherein the agent that alters activities of G protein coupled receptors and cAMP is selected from the group consisting of AB-MECA (*N*<sup>6</sup>-4-amino benzyl-5'-*N*-methylcarboxamidoadenosine), CPA (*N*<sup>6</sup>-cyclopentyladenosine), ADAC (*N*<sup>6</sup>-[4-[[[4-[[[(2-aminoethyl) amino] carbonyl] methyl]-anilino] carbonyl] methyl] phenyl] adenosine), CCPA (2-chloro-*N*<sup>6</sup>-cyclopentyladenosine), CHA (*N*<sup>6</sup>-cyclohexyladenosine), GR79236 (*N*<sup>6</sup>-[1*S*, *trans*,2-hydroxycyclopentyl] adenosine), *S*-ENBA ((2*S*)- *N*<sup>6</sup>-(2-endonorbanyl)adenosine), IAB-MECA (*N*<sup>6</sup>-(4-amino-3-iodobenzyl)adenosine-5'-*N*-methylcarboxamidoadenosine), *R*-PIA (*R*-*N*<sup>6</sup>-(phenylisopropyl) adenosine), ATL146e (4-{3-[6-amino-9-(5-ethylcarbamoyl-3,4-dihydroxy-tetrahydro-furan-2-yl)-9*H*-purin-2-yl]-prop-2-ynyl}-cyclohexanecarboxylic acid methyl ester), CGS-21680 (APEC or 2-[*p*-(2-carbonyl-ethyl)-phenyl ethyl amino]-5'-*N*-ethylcarboxamidoadenosine), CV1808 (2-phenylaminoadenosine), HNECA (2-hex-1-ynyl-5'-*N*-ethylcarboxamido adenosine), NECA (5'-*N*-ethyl-carboxamido adenosine),

PAPA-APEC (2-(4-[2-[(4-aminophenyl) methyl carbonyl]ethyl] phenyl) ethylamino-5'-N-ethyl carboxamidoadenosine), DITC-APEC (2-[*p*-(4-isothiocyanatophenylamino thiocarbonyl-2-ethyl)-phenylethylamino]-5'-N-ethylcarboxamidoadenosine), DPMA (*N*<sup>6</sup>-(2(3,5-dimethoxy phenyl)-2-(2-methyl phenyl) ethyl)adenosine), S-PHPNECA ((*S*)-2-phenylhydroxypropynyl-5'-N-ethylcarbox amidoadenosine), WRC-0470 (2-cyclohexylmethylidenehydrazinoadenosine), AMP-579 (1*S*-[1a,2b,3b,4a(S\*)]-4-[7-[[2-(3-chloro-2-thienyl)-1-methylpropyl]amino]-3H-imidazo [4,5-b] pyridyl-3-yl] cyclopentane carboxamide), IB-MECA (*N*<sup>6</sup>- (3-iodobenzyl) adenosine -5'-N-methyluronamide), 2-CIADO (2-chloroadenosine), I-ABA (*N*<sup>6</sup>-(4-amino-3-iodobenzyl) adenosine), *S*-PIA (*S*-*N*<sup>6</sup>-(phenylisopropyl)adenosine), 2-[(2-aminoethyl-aminocarbonylethyl) phenylethyl amino]-5'-N-ethyl-carboxamidoadenosine, 2-Cl-IB-MECA (2-chloro-*N*<sup>6</sup>- (3-iodobenzyl)adenosine-5'-N-methyluronamide), adenosine, polyadenylic acid, and any mixture thereof.